

*C' cont*

$R^2$  is an acyl group;

$CO_2R^3$  is a carboxy group or a carboxylate anion, or  $R^3$  is a readily removable carboxy protecting group;

$R^4$  represents hydrogen or up to four substituents selected from alkyl, alkenyl, alkynyl, alkoxy, hydroxy, halogen, amino, alkylamino, acylamino, dialkylamino,  $CO_2R$ ,  $CONR_2$ ,  $SO_2NR_2$  (where R is hydrogen or  $C_{1-6}$  alkyl), aryl and heterocyclyl, which may be the same or different and wherein any  $R^4$  alkyl substituent is optionally substituted by any other  $R^4$  substituent;

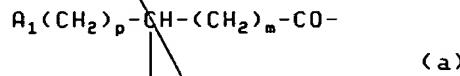
X is S, SO,  $SO_2$ , O or  $CH_2$ ;

m is 1 or 2;

n is 0;

"acyl" is selected from the group consisting of formula

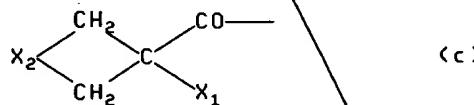
(a) to (f):



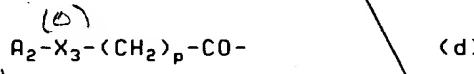
(a)



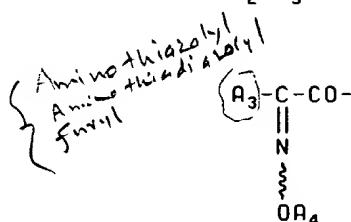
(b)



(c)



(d)



(e)



(f)

*C1*  
*Cont*

wherein p is 0, 1 or 2;

m is 0, 1 or 2;

*A<sub>1</sub>* is C<sub>1-6</sub> alkyl, substituted C<sub>1-6</sub> alkyl, C<sub>3-6</sub> cycloalkyl, cyclohexenyl, cyclohexadienyl, or an aromatic or heteroaromatic group;

*X<sub>1</sub>* is a hydrogen or halogen atom, a carboxylic acid, carboxylic ester, sulphonic acid, azido, tetrazolyl, hydroxy, acyloxy, amino, ureido, acylamino, heterocyclylamino, guanidino or acylureido group;

*A<sub>2</sub>* is an aromatic or heteroaromatic group, a substituted alkyl group, or a substituted dithietane;

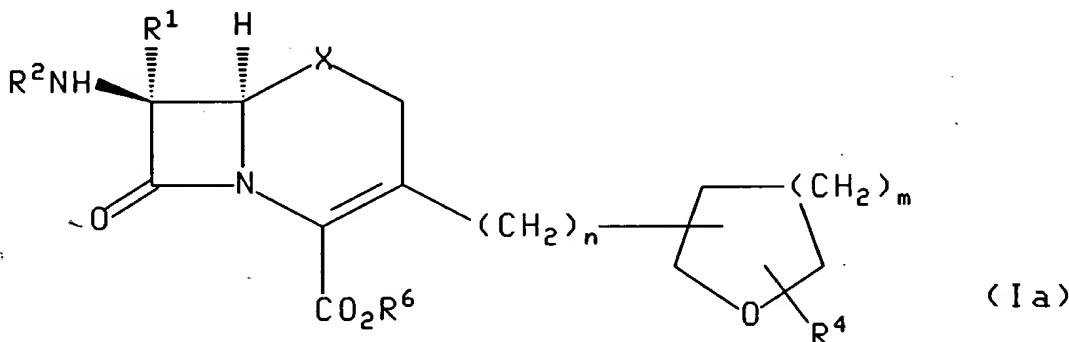
*X<sub>2</sub>* is a -CH<sub>2</sub>OCH<sub>2</sub>-, -CH<sub>2</sub>SCH<sub>2</sub>- or alkylene group;

*X<sub>3</sub>* is an oxygen or sulphur atom;

*A<sub>3</sub>* is an aryl or heteroaryl group; and

*A<sub>4</sub>* is hydrogen, C<sub>1-6</sub> alkyl, C<sub>3-8</sub> cycloalkyl, C<sub>3-8</sub> cycloalkyl(C<sub>1-6</sub>)alkyl, C<sub>1-6</sub> alkoxy carbonyl(C<sub>1-6</sub>)alkyl, C<sub>2-6</sub> alkenyl, carboxy(C<sub>1-6</sub>)alkyl, C<sub>2-6</sub> alkynyl, aryl or C<sub>1-6</sub> alkyl substituted by up to three aryl groups.

*58.* A compound as claimed in claim 57 having the formula (Ia):



wherein R<sup>1</sup>, R<sup>2</sup>, R<sup>4</sup>, m, n and X are as defined with respect to formula (I) in claim 57 and the group CO<sub>2</sub>R<sup>6</sup> is CO<sub>2</sub>R<sup>3</sup> where CO<sub>2</sub>R<sup>3</sup> is a carboxy group or a carboxylate anion, or a pharmaceutically acceptable salt or in vivo hydrolysable ester thereof.

*59.* A compound as claimed in claim 57 wherein R<sup>1</sup> is hydrogen.

4. A compound as claimed in claim ~~57~~ wherein  $A_1$  is optionally substituted phenyl,  $X_1$  is hydrogen or amino,  $A_2$  is optionally substituted phenyl,  $X_3$  is oxygen,  $A_3$  is aminothiazolyl, aminothiadiazolyl or furyl, and  $R^4$  is hydrogen,  $C_{1-6}$  alkyl, or carboxy  $C_{1-6}$  alkyl. /

5. A compound as claimed in claim ~~57~~ wherein  $CO_2R^3$  is carboxy or a carboxylate anion or  $R^3$  is  $t$ -butyl, 4-methoxybenzyl, diphenylmethyl, acetoxymethyl, acetoxyethyl, pivaloyloxymethyl, propan-2-yloxycarbonyloxyethyl or 2-ethoxycarbonyl-but-2-enyl. /

6. A compound as claimed in claim ~~57~~ wherein the cyclic ether group bonded to the 3-position of the cephalosporin nucleus is unsubstituted or substituted by up to three substituents selected from  $C_{1-6}$  alkyl,  $C_{1-6}$  alkoxy,  $C_{1-6}$  alkoxy carbonyl,  $C_{1-6}$  alkanoyloxy  $C_{1-6}$  alkyl or  $C_{1-6}$  alkoxy  $C_{1-6}$  alkyl. /

7. A compound as claimed in claim ~~57~~ wherein  $m$  is 1. /

8. A compound as claimed in claim ~~57~~ wherein the cyclic ether group is a tetrahydrofuran-2-yl or a tetrahydropyran-2-yl group. /

9. A compound as claimed in claim ~~57~~ selected from the group consisting of:

sodium (6R,7R)-7-[2-(2-aminothiazol-4-yl)-2-(Z)-methoxyiminoacetamido]-3-[(RS)-tetrahydrofuran-2-yl]ceph-3-em-4-carboxylate;

pivaloyloxymethyl (6R,7R)-7-[2-(2-aminothiazol-4-yl)-2-(Z)-methoxyiminoacetamido]-3-[(RS)-tetrahydrofuran-2-yl]ceph-3-em-4-carboxylate;

sodium (6R,7R)-7-[2-(2-aminothiazol-4-yl)-2-(Z)-methoxyiminoacetamido]-3-[(RS)-tetrahydropyran-2-yl]ceph-3-em-4-carboxylate;

pivaloyloxymethyl (6R,7R)-7-[2-(2-aminothiazol-4-yl)-2-(Z)-methoxyiminoacetamido]-3-[(RS)-tetrahydropyran-2-yl]ceph-3-em-4-carboxylate;

(6R, 7R)-7-[2-(2-aminothiazol-4-yl)-2-(Z)-hydroxyiminoacetamido]-3-[(RS)-tetrahydrofuran-2-yl]ceph-3-em-4-carboxylic acid;

sodium (6R, 7R)-7-[2-(2-aminothiazol-4-yl)-2-(Z)-methoxyiminoacetamido]-3-[(S)-tetrahydrofuran-2-yl]ceph-3-em-4-carboxylate;

pivaloyloxymethyl (6R, 7R)-7-[2-(2-aminothiazol-4-yl)-2-(Z)-methoxyiminoacetamido]-3-[(S)-tetrahydrofuran-2-yl]ceph-3-em-4-carboxylate;

sodium (6R, 7R)-7-[2-(2-aminothiazol-4-yl)-2-(Z)-methoxyiminoacetamido]-3-[(R)-tetrahydrofuran-2-yl]ceph-3-em-4-carboxylate;

pivaloyloxymethyl (6R, 7R)-7-[2-(2-aminothiazol-4-yl)-2-(Z)-methoxyiminoacetamido]-3-[(R)-tetrahydrofuran-2-yl]ceph-3-em-4-carboxylate;

diphenylmethyl (6R, 7R)-7-phenylacetamido-3-[(RS)-tetrahydrofuran-2-yl]ceph-3-em-4-carboxylate;

sodium (6R, 7R)-7-[2-(2-aminothiazol-4-yl)-2-(Z)-methoxyiminoacetamido]-3-[(RS)-tetrahydrofuran-3-yl]ceph-3-em-4-carboxylate;

acetoxyethyl (6R, 7R)-7-[2-(2-aminothiazol-4-yl)-2-(Z)-methoxyiminoacetamido]-3-[(S)-tetrahydrofuran-2-yl]ceph-3-em-4-carboxylate;

sodium (6R, 7R)-7-[2-(2-aminothiazol-4-yl)-2-(Z)-methoxyiminoacetamido]-3-(5-methoxymethyltetrahydrofuran-2-yl)ceph-3-em-4-carboxylate;

sodium (6R, 7R)-7-[2-(2-aminothiazol-4-yl)-(Z)-pent-2-enamido]-3-[(S)-tetrahydrofuran-2-yl]ceph-3-em-4-carboxylate;

sodium (6R, 7R)-7-[2-(2-aminothiazol-4-yl)-2-(Z)-methoxyiminoacetamido]-3-[(S)-tetrahydrofuran-2-yl]ceph-3-em-4-carboxylate;

(RS)-1-acetoxyethyl (6R, 7R)-7-[2-(2-aminothiazol-4-yl)-2-(Z)-methoxyiminoacetamido]-3-[(S)-tetrahydrofuran-2-yl]ceph-3-em-4-carboxylate;

(6R, 7R)-7-[2-(2-aminothiazol-4-yl)-2-(Z)-carboxymethoxyiminoacetamido]-3-[(RS)-tetrahydrofuran-2-yl]ceph-3-

em-4-carboxylic acid disodium salt;

sodium (6R,7R)-7-[(R)-2-amino-2-(4-hydrophenyl)-acetamido]-3-[(S)-tetrahydrofuran-2-yl]ceph-3-em-4-carboxylate;

sodium (1S,6R,7R)-7-[2-(2-aminothiazol-4-yl)-2-(Z)-methoxyiminoacetamido]-3-[(S)-tetrahydrofuran-2-yl]ceph-3-em-4-carboxylate-1-oxide;

sodium 7-[2-(2-aminothiazol-4-yl)-2-(Z)-methoxyiminoacetamido]-3-(tetrahydrofuran-2-yl)-1-carba-1-dethiaceph-3-em-4-carboxylate;

sodium (6R,7R)-7-[2-(2-aminothiazol-4-yl)-2-(Z)-methoxyiminoacetamido]-3-[(S)-tetrahydrofuran-2-yl]ceph-3-em-4-carboxylate-1,1-dioxide;

(RS)-1-(propan-2-yl)oxycarbonyloxyethyl (6R,7R)-7-[2-(2-aminothiazol-4-yl)-2-(Z)-methoxyiminoacetamido]-3-[(S)-tetrahydrofuran-2-yl]ceph-3-em-4-carboxylate;

sodium (6R,7R)-7-[2-(2-aminothiazol-4-yl)-2-(Z)-methoxyiminoacetamido]-3-[(5R,2SR)-5-methyltetrahydrofuran-2-yl]ceph-3-em-4-carboxylate;

sodium (6R,7R)-7-[2-(furan-2-yl)-2-(Z)-methoxyiminoacetamido]-3-[(S)-tetrahydrofuran-2-yl]ceph-3-em-4-carboxylate;

sodium (6R,7R)-7-[2-(2-aminothiazol-4-yl)-2-(Z)-methoxyiminoacetamido]-3-[(S)-5,5-dimethyltetrahydrofuran-2-yl]ceph-3-em-4-carboxylate;

sodium (6R,7R)-7-[2-(2-aminothiazol-4-yl)-2-(Z)-methoxyiminoacetamido]-3-(5)-methoxycarbonyltetrahydrofuran-2-yl)ceph-3-em-4-carboxylate;

sodium (6R,7R)-7-[2-(2-aminothiazol-4-yl)-2-(Z)-methoxyiminoacetamido]-3-[3-methyltetrahydrofuran-2-yl]ceph-3-em-4-carboxylate; and

2-ethoxycarbonyl-(Z)-but-2-enyl (6R,7R)-7-[2-(2-aminothiazol-4-yl)-2-(Z)-methoxyiminoacetamido]-3-[(S)-tetrahydrofuran-2-yl]ceph-3-em-4-carboxylate.

66. A pharmaceutical composition comprising a compound of claim 58 or a pharmaceutically acceptable salt or in vivo